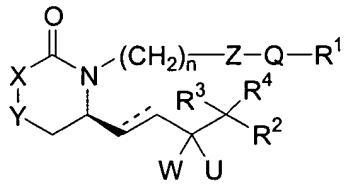
In the Claims

Amend the claims as follows:

1(Twice Amended). A compound having the structural formula I:



FORMULA I

or a pharmaceutically acceptable salt, enantiomer, diastereomer, prodrug or mixture thereof, wherein,

Q is $(CH_2)_m$, $(CH_2)_mC_{6-10}$aryl, $(CH_2)_mC_{5-10}$ heterocyclyl, $(CH_2)_mC_{3-10}$ heterocycloalkyl, $(CH_2)_mC_{3-8}$ cycloalkyl, $C(halo)_2$, said cycloalkyl, heterocycloalkyl, aryl or heterocyclyl unsubstituted or substituted with 1-3 groups of R^a;

X ~~[and Y independently]~~ represents ~~[CH_2,]~~ O, ~~[NR^9 or S, provided however, that X and Y are not O, NR^9 or S at the same time]~~;

Y represents CH_2;

U represents H, C1-3 alkyl or is not present when W is =O;

W represents OH or =O, provided that U is not present when W is =O;

R^1 represents $(CH_2)_p$hydroxy, $(CH_2)_pCN$, $(CH_2)_pCO_2R^{10}$, $(CH_2)_nSO_3R^6$, -$(CH_2)_pCF_2SO_2NH_2$, -$(CH_2)_pSO_2NH_2$, -$(CH_2)_pCONHSO_2R_2$, -$(CH_2)_pSO_2NHCOR^2$, -$(CH_2)_pPO(OH)_2$, $(CH_2)_pCONHPO_2R^6$, $(CH_2)_pCONHR^8$, $(CH_2)_pC_{1-4}$alkoxy, -$(CH_2)_p$cycloalkyl, ~~[$(CH_2)_p$-hydroxymethylketone]~~ $(CH_2)_pC(O)CH_2OH$ or $(CH_2)_n$heterocyclyl, said heterocyclyl unsubstituted or substituted with 1 to 3 groups of R^a and optionally containing an acidic hydroxyl group;

R^2 independently represents C_{1-10} alkyl, $(CH_2)_mC_{6-10}$aryl, $(CH_2)_mC_{5-10}$heterocyclyl, $(CH_2)_mC_{3-10}$ heterocycloalkyl, $(CH_2)_mC_{3-8}$ cycloalkyl, O-C_{1-10}alkyl, O-C_{6-10}aryl, O-C_{3-10}cycloalkyl, O-C_{3-10} heterocycloalkyl, O-C_{3-10} heterocycloalkyl, said alkyl, cycloalkyl, heterocycloalkyl, aryl or heterocyclyl unsubstituted or substituted with 1-3 groups of R^a;

R^3 and R^4 independently represents hydrogen, or C_{1-6} alkyl, or R^3 and R^4 may be taken together to form a 3-7 membered carbon ring optionally interrupted with 1-2 heteroatoms chosen from O, S, SO, SO_2, and NR^9;

R^6 and R^7 independently represents hydrogen, or C_{1-4} alkyl;

R^8 represents hydrogen, acyl, or sulfonyl;

R^9 represents hydrogen, C_{1-6} alkyl, said alkyl optionally substituted with 1-3 halogen, CN, OH, C_{1-6} alkoxy, C_{1-6} acyloxy or amino;

R^{10} represents hydrogen, C_{1-10} alkyl, C_{3-10} cyclcoalkyl, $(CH_2)_pC_{6-10}$ aryl, $(CH_2)_pC_{5-10}$ heterocyclyl, $CR^6R^7OC(O)O$ C_{3-10} cycloalkyl or $CR^6R^7OC(O)O$ C_{1-10} alkyl;

Z represents a ~~[triple bond]~~C≡C, O, S, $(C(R^b)_2)_n$, or C~~[h]~~H=CH;

R^b represents hydrogen, C1-6 alkyl or halogen;

R^a represents C_{1-6} alkoxy, C_{1-6} alkyl, CF_3, nitro, amino, cyano, C_{1-6} alkylamino, halogen, ~~[or Ra further represents for aryls and heterocyclyl,]~~ SC_{1-6}alkyl, SC_{6-10}aryl, SC_{5-10}heterocyclyl, CO_2R^6, OC_{6-10}aryl, OC_{5-10}heterocyclyl, CH_2OC_{1-6} alkyl, CH_2SC_{1-6} alkyl, CH_2Oaryl, CH_2Saryl;

--- represents a double or single bond;

p represents 0-3;

n represents 0-4; and

m represents 0-8.

2(Original). A compound in accordance with claim 1 wherein R^1 is $(CH_2)_pCN$, $(CH_2)_pCO_2R^{10}$, $-(CH_2)_pPO(OH)_2$, $(CH_2)_pCONHPO_2R^6$, $(CH_2)_pCONHR^8$, or $(CH_2)_n$heterocyclyl, said heterocyclyl unsubstituted or substituted with 1 to 3 groups of R^a and all other variables are as originally described.

3 (Original). A compound in accordance with claim 2 wherein Z is a bond or S, Y is CH_2 and X is O, S or CH_2.

4 (Twice Amended). A compound in accordance with claim 1 wherein R^1 is $(CH_2)_pCO_2R^{10}$, X is O, ~~[and]~~ Y are CH_2, Z is $(C(R^b)_2)_n$, Q is (CH_2)m, and R^2 is $(CH_2)_mC_{6-10}$aryl, said aryl unsubstituted or substituted with 1 to 3 groups of R^a.

5 (Original). A compound in accordance with claim 2 wherein R^1 is $(CH_2)_mC_{5-10}$heterocyclyl, U is H, or C_{1-3} alkyl, W is OH, Z is a bond or S, R^2 is $(CH_2)_mC_{6-10}$aryl, said aryl unsubstituted or substituted with 1 to 3 groups of R^a, said heterocyclyl unsubstituted or substituted with 1 to 3 groups of R^a and all other variables are as originally described.

6 (Twice Amended). A compound which is:

~~[7-{(2*R*)-2-[(3*R*)-3-hydroxy-4-phenylbutyl]-6-oxopiperidin-1-yl}heptanoic acid;]~~

7-{(4*S*)-4-[(3*R*)-3-hydroxy-4-phenylbutyl]-2-oxo-1,3-oxazinan-3-yl}heptanoic acid;

~~7-{(4*S*)-4-[(3*R*)-3-hydroxy-4-phenylbutyl]-2-oxo-1,3-thiazinan-3-yl}heptanoic acid;~~

~~5-(3-{(2*R*)-2-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-6-oxopiperidin-1-yl}propyl)thiophene-2-carboxylic acid;~~

5-(3-{(4*R*)-4-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-2-oxo-1,3-oxazinan-3-yl}propyl)thiophene-2-carboxylic acid;

~~5-(3-{(4*R*)-4-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-2-oxo-1,3-thiazinan-3-yl}propyl)thiophene-2-carboxylic acid;~~

~~isopropyl 5-(3-{(2*R*)-2-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-6-oxopiperidin-1-yl}propyl)thiophene-2-carboxylate;~~

isopropyl 5-(3-{(4*R*)-4-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-2-oxo-1,3-oxazinan-3-yl}propyl)thiophene-2-carboxylate;

~~isopropyl 5-(3-{(4*R*)-4-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-2-oxo-1,3-thiazinan-3-yl}propyl)thiophene-2-carboxylate;~~

~~2-(3-{(2*R*)-2-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-6-oxopiperidin-1-yl}propyl)-1,3-thiazole-5-carboxylic acid;~~

~~5-(3-{(2*R*)-2-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-6-oxopiperidin-1-yl}propyl)-1,3-thiazole-2-carboxylic acid;~~

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~~5-(3-{(2*R*)-2-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-6-oxopiperidin-1-yl}propyl)-1,3-oxazole-2-carboxylic acid;~~
~~2-(3-{(2*R*)-2-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-6-oxopiperidin-1-yl}propyl)-1,3-oxazole-5-carboxylic acid;~~
~~5-(3-{(2*R*)-2-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-6-oxopiperidin-1-yl}propyl)-1,2λ^5,5λ^5-oxadiazole-2-carboxylic acid;~~
~~5-(3-{(2*R*)-2-[(1*E*,3*S*)-3-hydroxy-4-phenylbut-1-enyl]-6-oxopiperidin-1-yl}propyl)-4*H*-1,2,4-triazole-3-carboxylic acid;~~
~~6-[(1E)-(3R)-3-hydroxy-4-phenyl-but-1-enyl]-1-[6-(1H-tetrazol-5-yl)-hexyl]-piperidin-2-one;~~
~~7-{[(1*E*)-(2*R*)-2-(3*S*)-3-hydroxy-4-phenyl-but-1-enyl]-6-oxo-piperdin-1-yl}heptanoic acid;~~
~~isopropyl 7-{[(1*E*)-(2*R*)-2-(3*S*)-3-hydroxy-4-phenyl-but-1-enyl]-6-oxo-piperdin-1-yl}heptanoate;~~
~~isopropyl 7-{(2*R*)-2-[(3*R*)-3-hydroxy-4-phenyl-butyl]-6-oxo-piperdin-1-yl}-heptanoate;~~
~~7-{[(2*R*)-2-(3*R*)-3-hydroxy-4-phenyl-butyl]-6-oxo-piperdin-1-yl}heptanoic acid;~~
~~methyl 5-{3-[(2*R*)-2-((1*E*)-(3*S*)-3-hydroxy-4-phenyl-but-1-enyl)-6-oxo-piperidin-1-yl]-propyl}-thiophene-2-carboxylate;~~
~~5-{3-[(2*R*)-2-((1*E*)-(3*S*)-3-hydroxy-4-phenyl-but-1-enyl)-6-oxo-piperidin-1-yl]-propyl}-thiophene-2-carboxylic acid;~~
~~5-{3-[(2*R*)-2-((3*S*)-3-hydroxy-4-phenyl-butyl)-6-oxo-piperidin-1-yl]-propyl}-thiophene-2-carboxylic acid;~~
~~isopropyl5-{3-[(2*R*)-2-((1*E*)-(3*S*)3-hydroxy-4-phenyl-but-1-enyl)-6-oxo-piperidin-1-yl]-propyl}-thiophene-2-carboxylate;~~
~~isopropyl 5-{3-[(2*R*)-2-((3*S*)-3-hydroxy-4-phenyl-butyl)-6-oxo-piperidin-1-yl]-propyl}-thiophene-2-carboxylate;~~
~~6-[(3*R*)-3-hydroxy-4-phenyl-butyl]-1-[6-(1*H*-tetrazol-5-yl)-hexyl]-piperidin-2-one;~~
~~isopropyl 7-{(2*R*)-2-[(1*E*)-4,4-difluoro-3-oxo-4-phenylbut-1-enyl]-6-oxopiperidin-1-yl}heptanoate;~~
~~methyl 5-{3-[(2*R*)-2-((1*E*)-(3*S*)-3-hydroxy-4-phenyl-but-1-enyl)-6-oxo-piperidin-1-yl]-propyl}-thiophene-2-carboxylate;~~
~~5-{3-[(2*R*)-2-((3*S*)-3-hydroxy-4-phenyl-butyl)-6-oxo-piperidin-1-yl]-propyl}-thiophene-2-carboxylic acid;~~
~~isopropyl 5-{3-[(2*R*)-2-((3*S*)-3-hydroxy-4-phenyl-butyl)-6-oxo-piperidin-1-yl]-propyl}-thiophene-2-carboxylate;~~
~~6-[(3*R*)-3-hydroxy-4-phenyl-butyl]-1-[6-(1*H*-tetrazol-5-yl)-hexyl]-piperidin-2-one;~~
or a pharmaceutically acceptable salt, enantiomer, diastereomer, prodrug or mixture thereof.

7. Canceled.

8 (Original). A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a compound of formula I, as recited in claim 1.

9 (Original). A method for treating ocular hypertension or glaucoma comprising administration to a patient in need of such treatment a therapeutically effective amount of a compound of claim 1, said compound administered in a topical formulation as a solution or suspension.

10 (Original). The method according to claim 9 wherein one or more active ingredients belonging to the group consisting of: β-adrenergic blocking agent, parasympatho-mimetic agent, sympathomimetic agent, carbonic anhydrase inhibitor, Maxi-K channel blocker, and a prostaglandin, hypotensive lipid, neuroprotectant, and 5-HT2 receptor agonist is added to the topical formulation.

11 (Original). The method according to claim 10 wherein the β-adrenergic blocking agent is timolol, betaxolol, levobetaxolol, carteolol, or levobunolol; the parasympathomimetic agent is pilocarpine; the sympathomimetic agent is epinephrine, brimonidine, iopidine, clonidine, or para-aminoclonidine, the carbonic anhydrase inhibitor is dorzolamide, acetazolamide, metazolamide or brinzolamide; COSOPT®, the Maxi-K is Penitrem A, paspalicine, charybdotoxin, iberiotoxin, Paxicillan, Aflitram, Verroculogen, 1-(1-isobutyl-6-methoxy-1H-indazol-3-yl)-2-methylpropan-1-one; 1-[1-(2,2-dimethylpropyl)-6-methoxy-1H-indazol-3-yl]-2-methylpropan-1-one; 1-[1-(cyclohexylmethyl)-6-methoxy-1H-indazol-3-yl]-2-methylpropan-1-one; 1-(1-hexyl-6-methoxy-1H-indazol-3-yl)-2-methylpropan-1-one; 1-[1-(2-ethylhexyl)-6-methoxy-1H-indazol-3-yl]-2-methylpropan-1-one; 1-(3-isobutyryl-6-methoxy-1H-indazol-1-yl)buan-2-one; 1-(3-isobutyryl-6-methoxy-1H-indazol-1-yl)-3,3-dimethylbutan-2-one; 1-(3-cyclopentylcarbonyl)-6-methoxy-1H-indazol-1-yl)-3,3-dimethylbutan-2-one; 1-(3,3-dimethyl-2-oxobutyl) -6-methoxy-1H-indazole-3-carboxylic acid; and 1-[3-(3-hydroxypropanoyl) -6-methoxy-1H-indazol-1-yl]-3,3-dimethylbutan-2-one, the prostaglandin is latanoprost, travaprost, unoprostone, rescula, or S1033, the hypotensive lipid is lumigan, the neuroprotectant is eliprodil, R-eliprodil or memantine; and the 5-HT2 receptor agonist is 1-(2-aminopropyl)-3-methyl-1H-imdazol-6-ol fumarate or 2-(3-chloro-6-methoxy-indazol-1-yl)-1-methyl-ethylamine.

12 (Original). A method for treating macular edema or macular degeneration, treating dry eye, increasing retinal and optic nerve head blood velocity, increasing retinal and optic nerve oxygen tension or providing a neuroprotection, comprising administration to a patient in need of such treatment a pharmaceutically effective amount of a compound of a compound as recited in claim 1.

13 (Original). The method according to claim 9 in which the topical formulation optionally contains xanthan gum or gellan gum.

14 (Original). A method for stimulating bone formation, treating or reducing the risk of contracting a disease state or condition related to abnormal bone

resorption, in a mammal in need thereof comprising administering to said mammal a therapeutically effective amount of a compound as recited in claim 1.

15 (Original). The method according to claim 14 wherein said disease state or condition is selected from the group consisting of osteoporosis, glucocorticoid induced osteoporosis, Paget's disease, abnormally increased bone turnover, periodontal disease, tooth loss, bone fractures, rheumatoid arthritis, periprosthetic osteolysis, osteogenesis imperfecta, metastatic bone disease, hypercalcemia of malignancy, and multiple myeloma.

16 (Original). The method according to claim 14 wherein a bisphosphonate active selected from the group consisting of alendronate, cimadronate, clodronate, tiludronate, etidronate, ibandronate, neridronate, olpandronate, risedronate, piridronate, pamidronate, zolendronate, pharmaceutically acceptable salts thereof, and mixtures thereof is optionally added.

17 (Original). The method according to Claim 16 comprising administering another agent selected from an organic bisphosphonate; a cathepsin K inhibitor, an estrogen, an estrogen receptor modulator, an androgen receptor modulator, an inhibitor of osteoclast proton ATPase, an inhibitor of HMG-CoA reductase, an integrin receptor antagonist, an osteoblast anabolic agent, calcitonin, vitamin D, a synthetic Vitamin D analogue, or a pharmaceutically acceptable salt or mixture thereof.